

On page 1, replace paragraph on lines 19-35 with the following paragraph.

The invention features a method for diagnosing a malignant neoplasm in a mammal by contacting a bodily fluid from the mammal with an antibody which binds to an human aspartyl (asparaginyl) beta-hydroxylase (HAAH) polypeptide under conditions sufficient to form an antigen- antibody complex and detecting the antigen-antibody complex (for the purposes of this specification, HAAH polypeptide refers to the amino acid sequence of SEQ ID NO:2 and HAAH cDNA refers to the nucleotide sequence of SEQ ID NO:3). Malignant neoplasms detected in this manner include those derived from endodermal tissue, e.g., colon cancer, breast cancer, pancreatic cancer, liver cancer, and cancer of the bile ducts. Neoplasms of the central nervous system (CNS) such as primary malignant CNS neoplasms of both neuronal and glial cell origin and metastatic CNS neoplasms are also detected. Patient derived tissue samples, e.g., biopsies of solid tumors, as well as bodily fluids such as a CNS- derived bodily fluid, blood, serum, urine, saliva, sputum, lung effusion, and ascites fluid, are contacted with an HAAH-specific antibody.

On page 6, replace paragraph on lines 5-16 with the following amended paragraph.

For example, a compound which inhibits HAAH hydroxylation is a polypeptide that binds a HAAH ligand but does not transduce an intracellular signal or a polypeptide which contains a mutation in the catalytic site of HAAH. Such a polypeptide contains an amino acid sequence that is at least 50% identical to a naturally-occurring HAAH amino acid sequence or a fragment thereof and which has the ability to inhibit HAAH hydroxylation of substrates containing an EGF-like repeat sequence. More preferably, the polypeptide contains an amino acid sequence that is at least 75%, more preferably at least 85%, more preferably at least 95% identical to SEQ ID NO:2.

On page 47, lines 1-12, replace Table 4 with the following amended Table.

Table 4: Overexpression of enzymatically active HAAH
indicates malignancy

| Cdna | # of foci ± S.D. ^b | NIH 3T3 clone | # of colonies ^e |
|--------------------------------------|----------------------------------|-----------------------|----------------------------|
| pcDNA3 (mock) | 6.0 ± 3.3 | pcDNA (mock) | 0.4 ± 0.5 |
| murine [H]AAH | 14.0 ± 2.9 | clone 18 ^d | 6.2 ± 2.9 |
| mutant murine [H]AAH ^a | 1.6 ± 1.0 | clone 16 ^e | 4.7 ± 6.5 |
| [human] HAAH | 32.0 ± 5.4 | | |
| v-scr | 98.0 ± 7.1 | | |

a. enzymatically inactive [H]AAH

b. P<0.01 compared to mock and mutant murine [H]AAH

c. P<0.001 compared to mock

d. Clone 18 is a stable cloned NIH 3T3 cell line that overexpresses human HAAH by approximately two fold.

e. Clone 16 is a stable cloned NIH 3T3 cell line that overexpresses human HAAH by about 50%.